		Docket No. SN 09/97	
	FAX COVER SHEET	<ul> <li>Urgent</li> <li>Confidential</li> <li>Action Required</li> <li>Reply Requested</li> <li>For Your Info.</li> </ul>	
TO:	Art Unit 1631		
LOCATION:	USPTO		OFFICIAL
Fax No.:	703-872-9306 (Before Final Facsim	ile No.)	RECEIVED
FROM:	Carol See for Kamrin T. MacKnight Patent Attorney	t	CENTRAL FAX CENT
LOCATION:	GENENCOR INTERNATIONAL, INC. Legal Department 925 Page Mill Road Palo Alto, CA 94304-1013 Tel: 650-846-5838 Fax: 650-845-6504	•	
DATE:	November 3, 2003		
NUMBER OF F	PAGES TO FOLLOW: 8 SENT BY:	cas	
Re: USSN 09/975	,139; Docket No. GC637-2		
Attachments: Trai	nsmittal Letter (1 page) in duplicate, and Resprenent (6 pages).	ponse to	
The original of thi	s facsimile will be sent to you via:		
🗌 Regular Mail	☐ Overnight Mail ☐ Hand Delivery ☒ Will	Not Be Sen	t
🔀 Please initial a	cknowledgment of receipt by return facsimile	·	
client privilege and is in recipient, you are hereb strictly prohibited. If you	contained in this facsimile message is confidential and may itended only for the use of the above named individual(s). If y notified that any dissemination, distributh n or copying of a have received this communication in error, please notify us f) and return the original transmission to us by mail.	you are not the this communication	intended iti n is

GC637-2 faxcvf-pto

**P**002

11:34

Date: November 3, 2003

PATENT Docket No. GC637-2

## IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In re Application of	}
Volker Schellenberger et al.	) Group Art Unit: 1631
Serial No.: 09/975,139	) Examiner: Mahatan, Channing
Filed: October 10, 2001	)
For: Information Rich Libraries	

## TRANSMITTAL LETTER

Commissioner for Patents P.O. Box 1450 Alexandria, VA 22313-1450

Sir:

In response to the Restriction Requirement dated October 2, 2003, enclosed please find the following document: Response to Restriction Requirement.

The Commissioner is hereby authorized to charge any fees under 37 C.F.R. §§ 1.16 and 1.17 that may be required by this paper, and to credit any overpayment, to Deposit Account No. 07-1048 (Docket No. GC637-2). A duplicate of this paper is enclosed.

Respectfully submitted,

Date: November 3, 2003

Kamrin T. MacKnight Registration No. 38,230

Genencor International, Inc. 925 Page Mill Road Palo Alto, CA 94304-1013

T |: 650-846-5838 Fax: 650-845-6504

GC637-2 T-RR

**D**004

I hereby certify that this correspondence is being sent by facsimile transmission in accordance with § 1.6(d) addressed to Art Unit 1631, Before Final Facsimile No. (703) 872-9306, Commissioner for Patents, Alexandria, VA 22313-1450 on the date shown below.

• • • • • •	
Date:	November 3, 2003

y: \_\_\_\_

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

and a see

PATENT Docket No. GC637-2

RECEIVED CENTRAL FAX CENTER

OFFICIAL

in	re	Application	ı of
----	----	-------------	------

Group Art Unit: 1631

Volker Schellenberger et al.

Examiner: Mahatan, Channing

Serial No.: 09/975,139

Examiner: Manatan,

Filed: October 10, 2001

For: Information Rich Libraries

# Response to Restriction Requirement Mailed October 2, 2003

Commissioner for Patents P.O. Box 1450 Alexandria, VA 22313-1450

Sir:

In response to the Restriction Requirement mailed October 2, 2003, Applicants respectfully request that the following amendments be made. A complete list of the Claims, including marked-up versions of the rewritten, added, withdrawn, and/or cancelled claims is provided below, beginning on page 2. None of the amendments to the Claims is intended to narrow the scope of any of the amended Claims within the meaning of Festo<sup>1</sup>. The Remarks begin on page 6.

<sup>&</sup>lt;sup>1</sup> Festo Corp. v. Shoketsu Kogyo Kabushiki Co., No. 95-1066, 2000 WL 1753646 (Fed. Cir. Nov. 29, 2000).

#### LIST OF CLAIMS, SHOWING THE STATUS OF EACH CLAIM

Underlining denotes added text while strikethrough denotes deleted text.

### **IN THE CLAIMS:**

- 1. (Original) A method of creating a library of DNA sequences, said method comprising:
  - a) providing a DNA sequence that encodes a protein of interest;
  - b) providing a probability matrix for the protein;
  - c) providing a constraint vector for the protein;
- d) applying the constraint vector to the probability matrix to produce a substitution scheme recommending substitutions at at least two residues in the protein; and
- e) creating a library of DNA sequences incorporating changes in the DNA sequence that produce the recommended substitutions.
- 2. (Original) The method of claim 1, wherein said protein is selected from the group consisting of an esterase, dehydrogenase and hydrolase.
- 3. (Original) The method of claim 2, wherein said protein is selected from the group consisting of a protease, cellulase, lipase, hemicellulase, laccase, and amylase.
- 4. (Original) The method of claim 1, wherein said protein is selected from the group consisting of a transcription factor, growth factor, antibody, interleukin, antigen, and receptor.
- 5. (Original) The method of claim 1, wherein the probability matrix is based on structural characteristics selected from the group consisting of conservative residues, sequence alignments, three dlm national structure, residue environment, solvent accessibility, residue chemistry, propensity for a particular secondary structure, and combinations thereof.

USSN 09/975,139

11:34

- 6. (Original) The method of claim 1, wherein the constraint vector is based on structural characteristics known to affect protein function selected from the group consisting of proximity to the site of functionality, distance of  $\alpha$  or  $\beta$  carbons, contact with residues of interest, and contact with residues that contact the residue of interest.
- 7. (Original) The library of claim 1, wherein said library is a phage library.
  - 8. (Cancelled)
  - 9. (Cancelled)
  - 10. (Cancelled)
- 11. A system for creating libraries of nucleic acid sequences that encode variants of a protein, said system comprising:
  - an initial nucleic acid sequence that encodes a desired protein; a)
  - b) a probability matrix; and
  - c) a constraint vector.
  - 12. (Cancelled)
  - 13. (Cancelled)
- 14. (Original) The method of claim 1, wherein a library comprising at least 25 unique DNA sequences is produced.
- 15. (Original) The method of claim 14, wherein a library comprising at least 100 unique DNA sequences is produced.
- 16. (Original) The method of claim 15, wherein a library comprising at least 250 unique DNA sequences is produced.

- 17. (Original) The method of claim 16, wherein a library comprising at least 1000 unique DNA sequences is produced.
- 18. (Original) The method of claim 17, wherein a library comprising at least 2500 unique DNA sequences is produced.
- 19. (Original) The method of claim 18, wherein a library comprising at least 10,000 unique DNA sequences is produced.
- 20. (Original) The method of claim 1, wherein a library of less than 10<sup>9</sup> unique DNA sequences is produced.
- 21. (Original) The method of claim 20, wherein a library of less than 10<sup>6</sup> unique DNA sequences is produced.
- 22. (Original) The method of claim 21, wherein a library of less than 10<sup>5</sup> unique DNA sequences is produced
- 23. (Original) The method of claim 1, wherein the probability matrix is an algorithm.
- 24. (Original) The method of claim 1, wherein the probability matrix is generated by a computer.
- 25. (Original) The method of claim 1, wherein the constraint vector is an algorithm.
- 26. (Original) The method of claim 1, wherein the constraint vector is generated by a computer.
- 27. (Original) The method of claim 1, wherein the constraint vector is applied to the probability matrix using a computer.
  - 28. (Original) The method of claim 1, wherein the probability matrix is

USSN 09/975,139

normalized.

- 29. (Original) The method of claim 1, wherein the DNA sequence is generated from DNA shuffling.
  - 30. (Cancelled)
- 31. (Original) A method of creating a library of DNA sequences, said method comprising:
- a) providing a substitution scheme produced by applying a constraint vector to a probability matrix wherein the substitution scheme recommends substitutions at at least two residues in a protein of interest; and
- b) creating a library of DNA sequences incorporating substitutions in a DNA sequence encoding the protein of interest to create a library comprising the recommended substitutions.

USSN 09/975,139

#### REMARKS

The present application was originally filed with 31 Claims. In the present Restriction Requirement, the Examiner has restricted the Claims into five Groups:

- Group I contains Claims 1-6, 11, 14-29, and 31, directed to methods and systems for creating libraries of nucleic acid sequences;
  - Group II contains Claim 7, directed to a library;
- 3) Group III contains Claims 8, 9, and 30, directed to methods for screening libraries for a protein with an increase in a property of interest;
  - 4) Group IV contains Claim 10, directed to a protein; and
- 5) Group V contains Claims 12 and 13, directed to methods for improving a desired parameter of a protein of interest.

The Examiner argues that the Groups represent separate and patentably distinct inventions. While Applicants must respectfully traverse the restriction requirement, Applicants hereby elects the Claims in Group I (Claims 1-6, 11, 14-29, and 31), directed to methods and systems for creating libraries of nucleic acid sequences. Claims 8-10, 12, 13 and 30 have been cancelled. As Claim 7 (Group II) is a product-by-process Claim that is dependent upon Claim 1, Applicants respectfully request that this Group be joined with Group I. As Claim 7 is directed toward a particular type of library (*i.e.*, a phage library), Applicants submit that this is merely one type of library that is encompassed by Claim 1. Thus, there is no undue search burden to search Claims 1-7, 11, 14-29, and 31 together.

Applicants reserve the right to file Divisional application(s) to pursue the Claims cancelled herein. Should the Examiner have any questions regarding this application, she is encouraged to call the undersigned.

Respectfully submitted,

Kamrin T. MacKnight Registration No. 38,230

Date: November 3, 2003

Genencor International, Inc. 925 Page Mill Road
Palo Alto, CA 94304-1013

Tel: 650 846-5838 Fax: 650 845-6504